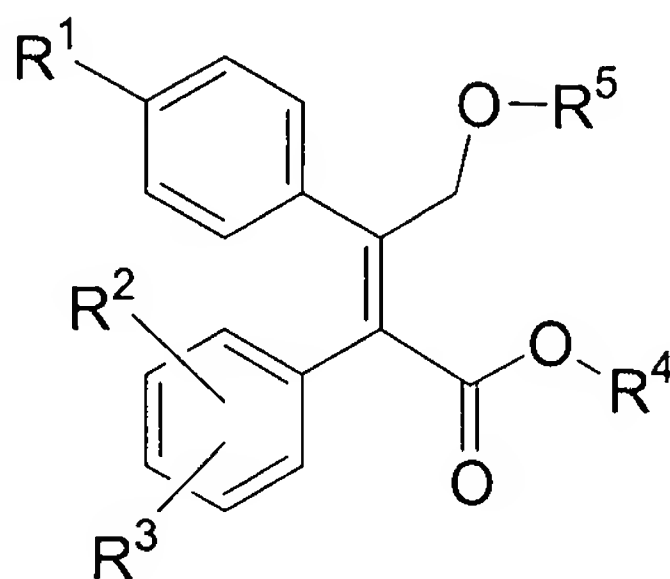


**Amendments to the Claims:**

This listing of claims will replace all prior versions, and listings, of claims in the application:

**Listing of Claims:**

1. (original) A compound of Formula I



I

or a pharmaceutically acceptable salt thereof, wherein

R<sup>1</sup> is selected from the group consisting of:

- (a) S(O)<sub>2</sub>CH<sub>3</sub>,
- (b) S(O)<sub>2</sub>NH<sub>2</sub>,
- (c) S(O)<sub>2</sub>NHC(O)CF<sub>3</sub>,
- (d) S(O)(NH)CH<sub>3</sub>,
- (e) S(O)(NH)NH<sub>2</sub>,
- (f) S(O)(NH)NHC(O)CF<sub>3</sub>,
- (g) P(O)(CH<sub>3</sub>)OH, and
- (h) P(O)(CH<sub>3</sub>)NH<sub>2</sub>;

R<sup>2</sup> and R<sup>3</sup> each are independently selected from the group consisting of:

- (a) hydrogen,
- (b) halo,
- (c) C<sub>1</sub>-6alkoxy,
- (d) C<sub>1</sub>-6alkylthio,
- (e) CN,
- (f) CF<sub>3</sub>,
- (g) C<sub>1</sub>-6alkyl, and

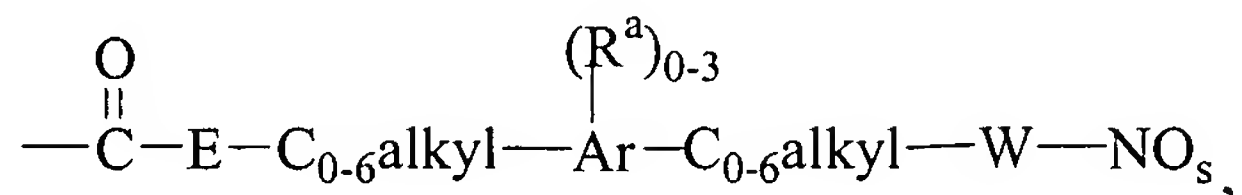
(h) N<sub>3</sub>;

R<sup>4</sup> is selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1-6</sub>alkyl, optionally substituted with 1-3 substituents independently selected from the group consisting of:
  - (i) halo,
  - (ii) phenyl, naphthyl or HET<sup>1</sup>, each of said phenyl, naphthyl or HET<sup>1</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>,
  - (iii) N(R<sup>i</sup>)R<sup>ii</sup>, wherein R<sup>i</sup> and R<sup>ii</sup> are each independently selected from the group consisting of hydrogen and C<sub>1-4</sub>alkyl,
  - (iv) -CO<sub>2</sub>R<sup>iii</sup>, wherein R<sup>iii</sup> is hydrogen or C<sub>1-4</sub>alkyl,
- (c) phenyl, naphthyl or HET<sup>2</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>5</sup> is selected from the group consisting of:

- (a) -NO<sub>s</sub>,
- (b) -C(O)-E-C<sub>1-10</sub>alkyl-W-NO<sub>s</sub>,
- (c)



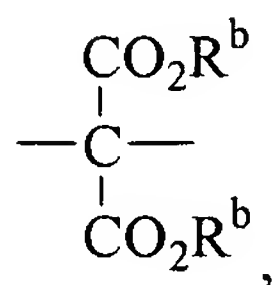
wherein:

each s is independently 1 or 2,

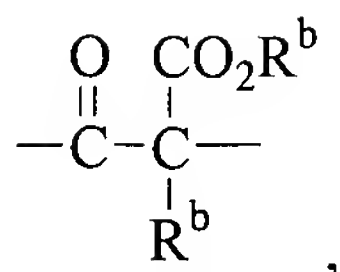
E is a bond, oxygen, sulfur or -C(O)-O-,

each W is independently selected from the group consisting of:

- (1) oxygen,
- (2) sulfur,
- (3)



- (4)



Ar is selected from the group consisting of: phenyl, naphthyl and HET<sup>3</sup>,

each R<sup>a</sup> is independently selected from the group consisting of:

- (1) halo,
- (2) C<sub>1-6</sub>alkyl,
- (3) C<sub>1-6</sub>alkoxy,
- (4) C<sub>1-6</sub>alkylthio,
- (5) OH,
- (6) CN,
- (7) CF<sub>3</sub>,
- (8) CO<sub>2</sub>R<sup>7</sup>, and
- (9) C<sub>0-6</sub>alkyl-W-NO<sub>s</sub>;

each R<sup>b</sup> is independently selected from the group consisting of:

- (1) C<sub>1-6</sub>alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>4</sup>, each of said phenyl, naphthyl or HET<sup>4</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>; and
- (2) phenyl, naphthyl or HET<sup>5</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>;

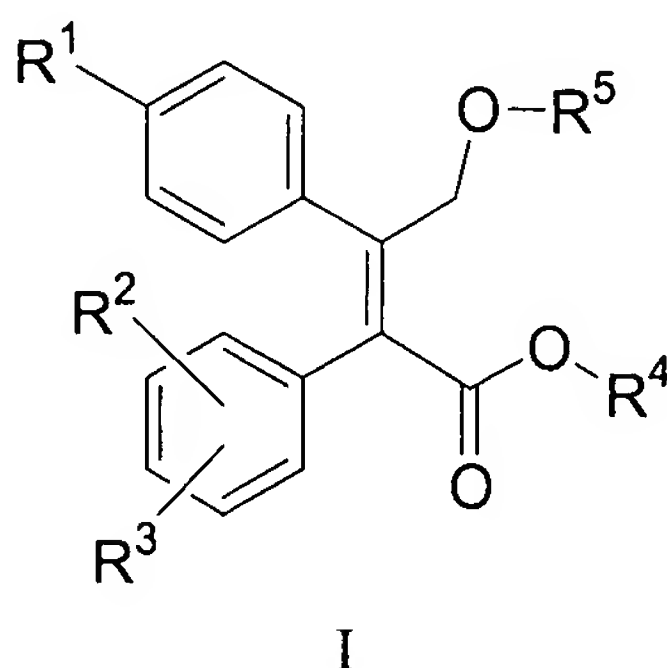
R<sup>6</sup>, R<sup>7</sup> and R<sup>8</sup> are each independently selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1-6</sub>alkyl; and

HET<sup>1</sup>, HET<sup>2</sup>, HET<sup>3</sup>, HET<sup>4</sup> and HET<sup>5</sup> are each independently selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolinyl, furanyl, imidazolyl, indolinyl, indolyl, indolazinyl, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidyl, 1,4-dioxanyl,

hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidyl, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

2. (original) A compound according to Claim 1 of Formula I



or a pharmaceutically acceptable salt thereof, wherein

R<sup>1</sup> is selected from the group consisting of:

- (a) S(O)<sub>2</sub>CH<sub>3</sub>,
- (b) S(O)<sub>2</sub>NH<sub>2</sub>,
- (c) S(O)<sub>2</sub>NHC(O)CF<sub>3</sub>,
- (d) S(O)(NH)CH<sub>3</sub>,
- (e) S(O)(NH)NH<sub>2</sub>,
- (f) S(O)(NH)NHC(O)CF<sub>3</sub>,
- (g) P(O)(CH<sub>3</sub>)OH, and
- (h) P(O)(CH<sub>3</sub>)NH<sub>2</sub>;

R<sup>2</sup> and R<sup>3</sup> each are independently selected from the group consisting of:

- (a) hydrogen,
- (b) halo,
- (c) C<sub>1</sub>-6alkoxy,
- (d) C<sub>1</sub>-6alkylthio,
- (e) CN,

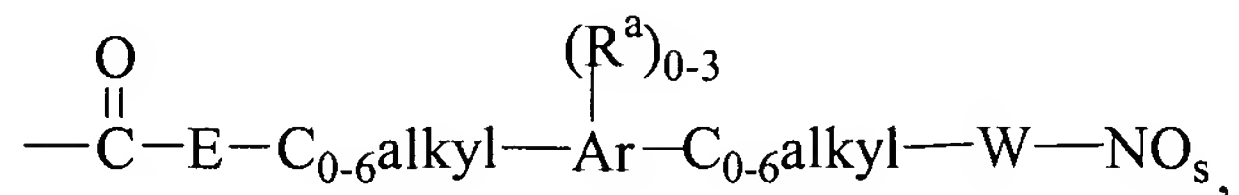
- (f) CF<sub>3</sub>,  
(g) C<sub>1-6</sub>alkyl, and  
(h) N<sub>3</sub>;

$R^4$  is selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1-6</sub>alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>1</sup>, each of said phenyl, naphthyl or HET<sup>1</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;
- (c) phenyl, naphthyl or HET<sup>2</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>5</sup> is selected from the group consisting of:

- (a) -NO<sub>2</sub>,  
(b) -C(O)-E-C<sub>1-10</sub>alkyl-W-NO<sub>2</sub>,  
(c)



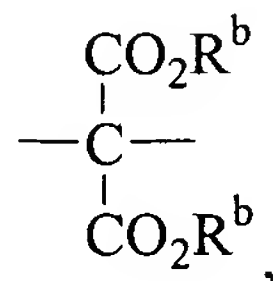
wherein:

each  $s$  is independently 1 or 2,

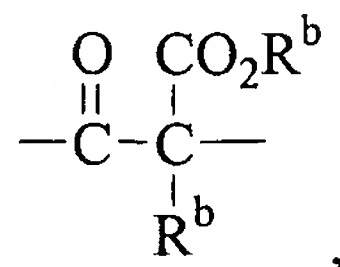
E is a bond, oxygen, sulfur or  $-C(O)-O-$ ,

each W is independently selected from the group consisting of:

- (1) oxygen,
- (2) sulfur,
- (3)



- (4)



Ar is selected from the group consisting of: phenyl, naphthyl and HET<sup>3</sup>,

each R<sup>a</sup> is independently selected from the group consisting of:

- (1) halo,
- (2) C<sub>1-6</sub>alkyl,
- (3) C<sub>1-6</sub>alkoxy,
- (4) C<sub>1-6</sub>alkylthio,
- (5) OH,
- (6) CN,
- (7) CF<sub>3</sub>,
- (8) CO<sub>2</sub>R<sup>7</sup>, and
- (9) C<sub>0-6</sub>alkyl-W-NO<sub>2</sub>;

each R<sup>b</sup> is independently selected from the group consisting of:

- (1) C<sub>1-6</sub>alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>4</sup>, each of said phenyl, naphthyl or HET<sup>4</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>; and
- (2) phenyl, naphthyl or HET<sup>5</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>;

R<sup>6</sup>, R<sup>7</sup> and R<sup>8</sup> are each independently selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1-6</sub>alkyl; and

HET<sup>1</sup>, HET<sup>2</sup>, HET<sup>3</sup>, HET<sup>4</sup> and HET<sup>5</sup> are each independently selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolinyl, furanyl, imidazolyl, indolinyl, indolyl, indolaziny, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidiny, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl,

dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidiny, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

3. (original) The compound according to Claim 2 wherein

R<sup>1</sup> is S(O)<sub>2</sub>CH<sub>3</sub>, and

R<sup>2</sup> and R<sup>3</sup> are both hydrogen.

4. (original) The compound according to Claim 3 wherein:

R<sup>4</sup> is C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>1</sup>, each of said phenyl, naphthyl or HET<sup>1</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>6</sup> is selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1</sub>-6alkyl; and

HET<sup>1</sup> is selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolinyl, furanyl, imidazolyl, indolinyl, indolyl, indolaziny, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidiny, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidiny, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

5. (original) The compound according to Claim 4 wherein R<sup>4</sup> is methyl, ethyl, propyl or isopropyl.

6. (original) The compound according to Claim 3 wherein:

R<sup>4</sup> is phenyl, naphthyl or HET<sup>2</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>6</sup> is selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1</sub>-6alkyl; and

HET<sup>2</sup> is selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolinyl, furanyl, imidazolyl, indolinyl, indolyl, indolaziny, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidyl, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidyl, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

7. (original) The compound according to Claim 3 wherein R<sup>5</sup> is -NO<sub>s</sub>, wherein s is 1 or 2.

8. (original) The compound according to Claim 3 wherein R<sup>5</sup> is -C(O)-E-C<sub>1</sub>-10alkyl-W-NO<sub>s</sub>, wherein:

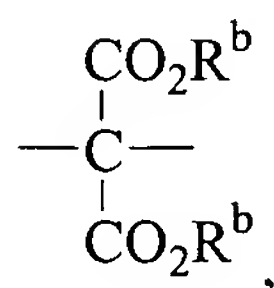
s is 1 or 2,



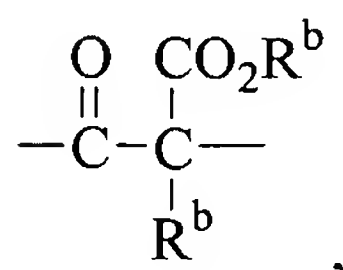
E is a bond, oxygen, sulfur or  $-C(O)-O-$ ,

W is selected from the group consisting of:

- (1) oxygen,
- (2) sulfur,
- (3)



- (4)



each  $\text{R}^b$  is independently selected from the group consisting of:

- (1)  $\text{C}_{1-6}$ alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or  $\text{HET}^4$ , each of said phenyl, naphthyl or  $\text{HET}^4$  being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo,  $\text{C}_{1-6}$ alkyl,  $\text{C}_{1-6}$ alkoxy,  $\text{C}_{1-6}$ alkylthio, OH, CN,  $\text{CF}_3$ , and  $\text{CO}_2\text{R}^8$ ; and
- (2) phenyl, naphthyl or  $\text{HET}^5$ , each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo,  $\text{C}_{1-6}$ alkyl,  $\text{C}_{1-6}$ alkoxy,  $\text{C}_{1-6}$ alkylthio, OH, CN,  $\text{CF}_3$ , and  $\text{CO}_2\text{R}^8$ ;

$\text{R}^8$  is selected from the group consisting of

- (a) hydrogen and
- (b)  $\text{C}_{1-6}$ alkyl; and

$\text{HET}^4$  and  $\text{HET}^5$  are each independently selected from the group consisting of:

benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolyl, furanyl, imidazolyl, indolyl, indolaziny, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidyl, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl,

dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidiny, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

9. (original) The compound according to Claim 8 wherein:

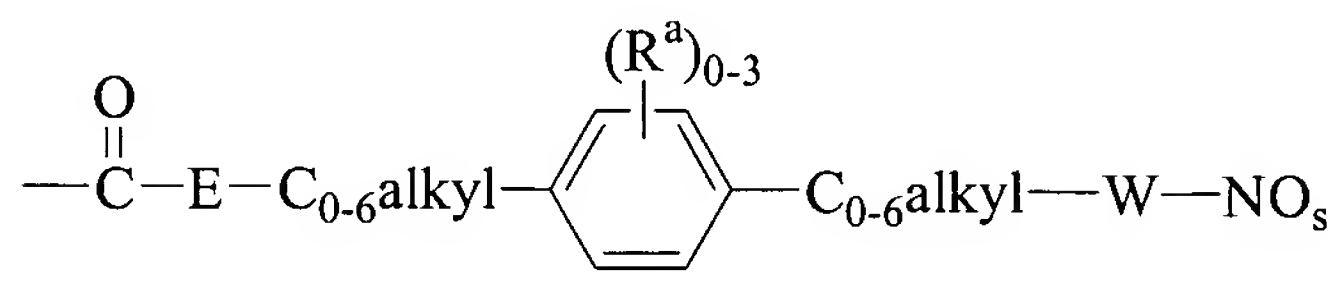
E is a bond or oxygen;

s is 2;

W is oxygen; and

R<sup>4</sup> is hydrogen, methyl, ethyl, propyl or isopropyl.

10. (original) The compound according to Claim 3 wherein R<sup>5</sup> is



wherein:

each s independently 1 or 2,

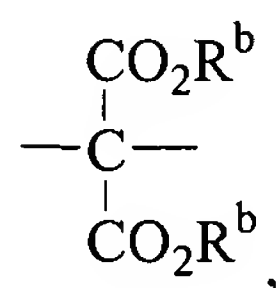
E is a bond, oxygen, sulfur or —C(O)—O—,

each W is independently selected from the group consisting of:

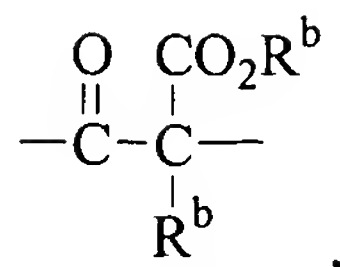
(1) oxygen,

(2) sulfur,

(3)



(4)



each R<sup>a</sup> is independently selected from the group consisting of:

- (1) halo,
- (2) C<sub>1</sub>-6alkyl,
- (3) C<sub>1</sub>-6alkoxy,
- (4) C<sub>1</sub>-6alkylthio,
- (5) OH,
- (6) CN,
- (7) CF<sub>3</sub>,
- (8) CO<sub>2</sub>R<sup>7</sup>, and
- (9) C<sub>0</sub>-6alkyl-W-NO<sub>s</sub>;

each R<sup>b</sup> is independently selected from the group consisting of:

- (1) C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>4</sup>, each of said phenyl, naphthyl or HET<sup>4</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>; and
- (2) phenyl, naphthyl or HET<sup>5</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>;

R<sup>7</sup> and R<sup>8</sup> is selected from the group consisting of

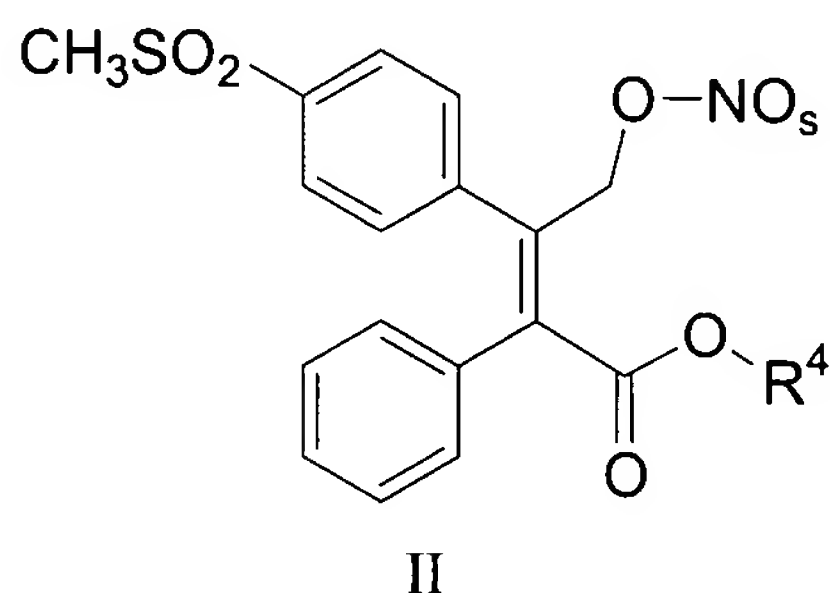
- (a) hydrogen and
- (b) C<sub>1</sub>-6alkyl; and

HET<sup>4</sup> and HET<sup>5</sup> are each independently selected from the group consisting of:

benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolinyl, furanyl, imidazolyl, indolinyl, indolyl, indolaziny, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidiny, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl,

pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidyl, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

11. (original) A compound according to Claim 2 of Formula II



or a pharmaceutically acceptable salt thereof, wherein

R<sup>4</sup> is selected from the group consisting of:

- (a) C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>1</sup>, each of said phenyl, naphthyl or HET<sup>1</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;
- (b) phenyl, naphthyl or HET<sup>2</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>6</sup> is selected from the group consisting of

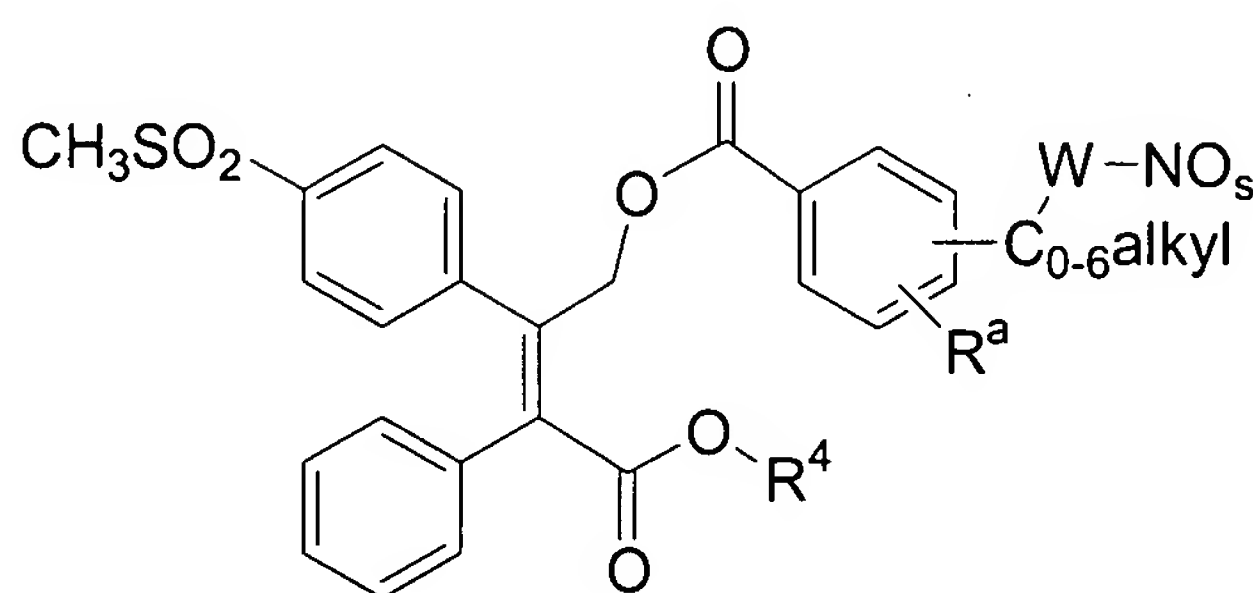
- (a) hydrogen and
- (b) C<sub>1</sub>-6alkyl;

s is 1 or 2; and

HET<sup>1</sup> and HET<sup>2</sup> are each independently selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolyl, furanyl, imidazolyl, indolyl, indolyl, indolaziny, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidyl, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidyl, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

12. - 14. (canceled)

15. (original) A compound according to Claim 2 of Formula III



III

or a pharmaceutically acceptable salt thereof, wherein

R<sup>4</sup> is selected from the group consisting of:

- (a) C<sub>1-6</sub>alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>1</sup>, each of said phenyl, naphthyl or

HET<sup>1</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

- (b) phenyl, naphthyl or HET<sup>2</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>6</sup> is selected from the group consisting of

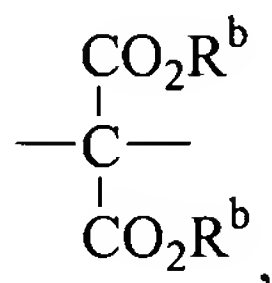
- (a) hydrogen,  
(b) C<sub>1</sub>-6alkyl;

R<sup>a</sup> is hydrogen or C<sub>0</sub>-6alkyl-W-NO<sub>s</sub>.

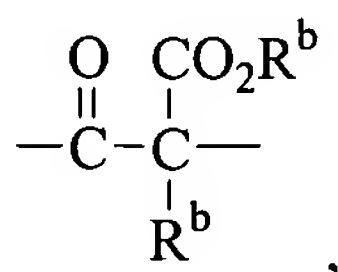
each s is independently 1 or 2,

each W is independently selected from the group consisting of:

- (1) oxygen,  
(2) sulfur,  
(3)



- (4)



each R<sup>b</sup> is independently selected from the group consisting of:

- (1) C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>4</sup>, each of said phenyl, naphthyl or HET<sup>4</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>; and  
(2) phenyl, naphthyl or HET<sup>5</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>;

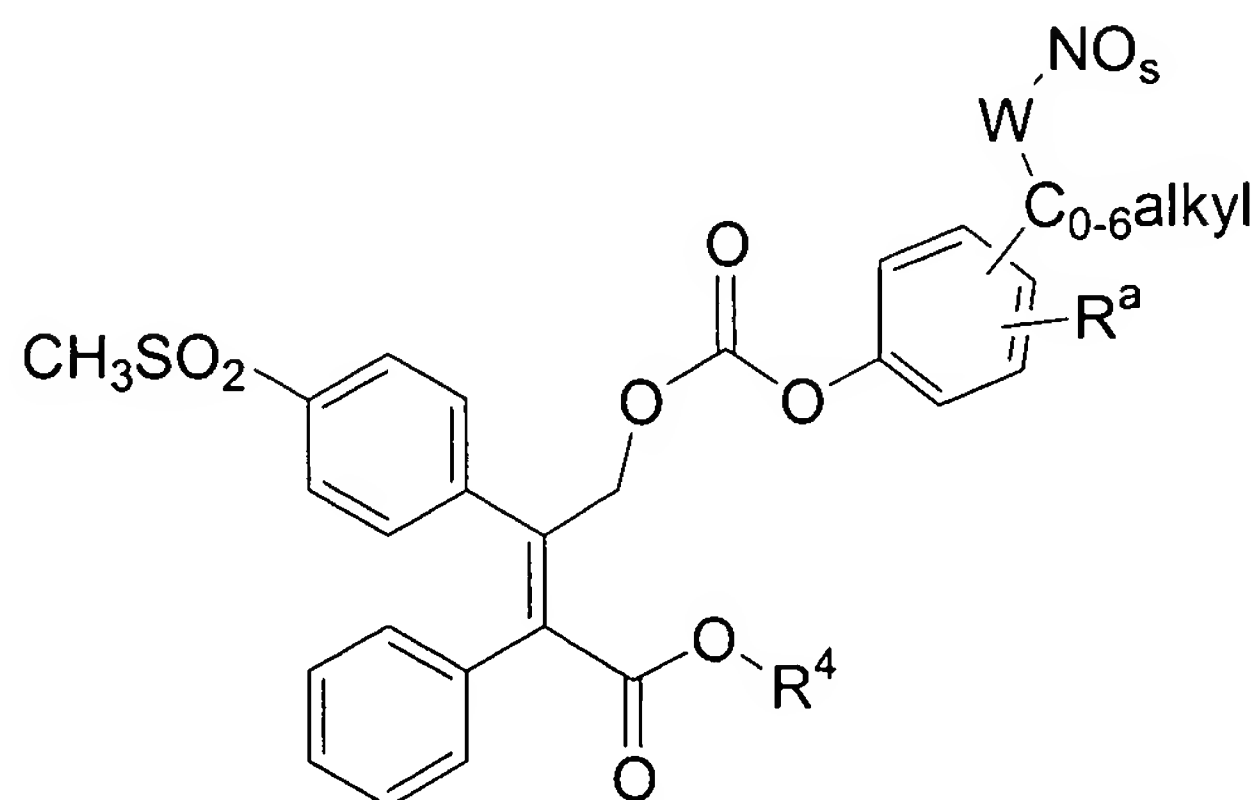
R<sup>8</sup> is selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1</sub>-6alkyl; and

HET<sup>1</sup>, HET<sup>2</sup>, HET<sup>4</sup> and HET<sup>5</sup> are each independently selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnoliny, furanyl, imidazolyl, indoliny, indolyl, indolaziny, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidiny, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinoliny, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidiny, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

16.-19. (canceled)

20. (original) A compound according to Claim 2 of Formula IV



IV

or a pharmaceutically acceptable salt thereof, wherein

R<sup>4</sup> is selected from the group consisting of:

- (a) C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>1</sup>, each of said phenyl, naphthyl or HET<sup>1</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;
- (b) phenyl, naphthyl or HET<sup>2</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>6</sup> is selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1</sub>-6alkyl;

R<sup>a</sup> is hydrogen or C<sub>0</sub>-6alkyl-W-NO<sub>s</sub>.

each s is independently 1 or 2;

each W is independently selected from the group consisting of:

- (1) oxygen,
- (2) sulfur,
- (3)
$$\begin{array}{c} \text{CO}_2\text{R}^b \\ | \\ -\text{C}- \\ | \\ \text{CO}_2\text{R}^b \end{array},$$
- (4)
$$\begin{array}{c} \text{O} \quad \text{CO}_2\text{R}^b \\ || \quad | \\ -\text{C}-\text{C}- \\ | \\ \text{R}^b \end{array},$$

each R<sup>b</sup> is independently selected from the group consisting of:

- (1) C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>4</sup>, each of said phenyl, naphthyl or HET<sup>4</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>; and



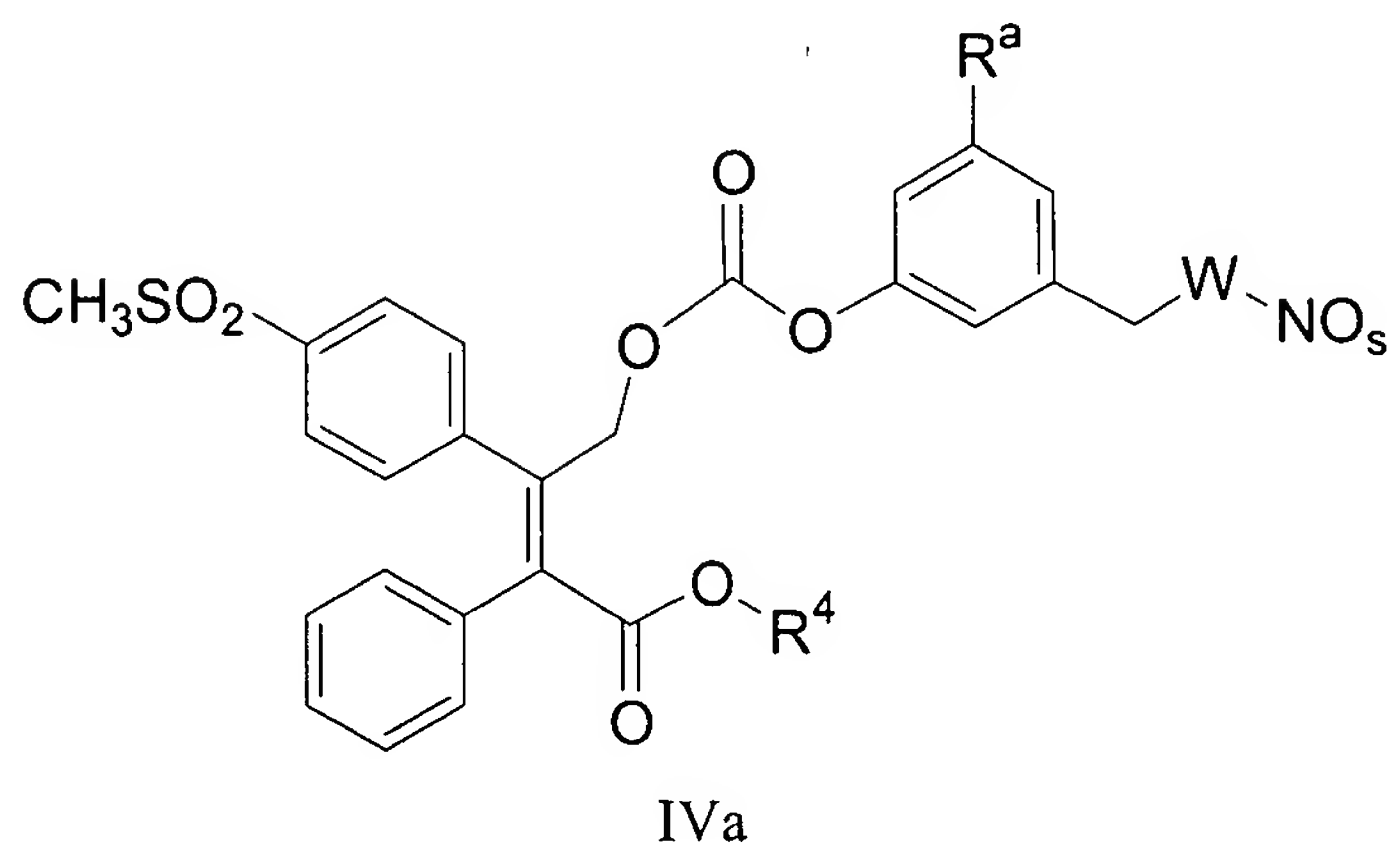
- (2) phenyl, naphthyl or HET<sup>5</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>;

R<sup>8</sup> is selected from the group consisting of

- (a) hydrogen,  
(b) C<sub>1</sub>-6alkyl; and

HET<sup>1</sup>, HET<sup>2</sup>, HET<sup>4</sup> and HET<sup>5</sup> are each independently selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolinyl, furanyl, imidazolyl, indolinyl, indolyl, indolazinyl, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidyl, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidyl, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

21. (original) The compound according to Claim 20 of Formula IVa



or a pharmaceutically acceptable salt thereof, wherein

R<sup>4</sup> is selected from the group consisting of:

- (a) C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>1</sup>, each of said phenyl, naphthyl or HET<sup>1</sup> being optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;
- (b) phenyl, naphthyl or HET<sup>2</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>6</sup>;

R<sup>6</sup> is selected from the group consisting of

- (a) hydrogen,
- (b) C<sub>1</sub>-6alkyl;

R<sup>a</sup> is hydrogen or C<sub>0</sub>-6alkyl-W-NO<sub>s</sub>.

each s is independently 1 or 2;

each W is independently selected from the group consisting of:

- (1) oxygen,
- (2) sulfur,
- (3)
$$\begin{array}{c} \text{CO}_2\text{R}^b \\ | \\ -\text{C}- \\ | \\ \text{CO}_2\text{R}^b, \end{array}$$

- (4)
$$\begin{array}{c} \text{O} \quad \text{CO}_2\text{R}^b \\ || \quad | \\ -\text{C}-\text{C}- \\ | \\ \text{R}^b, \end{array}$$

each R<sup>b</sup> is independently selected from the group consisting of:

- (1) C<sub>1</sub>-6alkyl, optionally substituted with 1-3 halo groups or optionally substituted with phenyl, naphthyl or HET<sup>4</sup>, each of said phenyl, naphthyl or HET<sup>4</sup> being optionally substituted with 1-3 substituents independently

selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>; and

- (2) phenyl, naphthyl or HET<sup>5</sup>, each optionally substituted with 1-3 substituents independently selected from the group consisting of: halo, C<sub>1</sub>-6alkyl, C<sub>1</sub>-6alkoxy, C<sub>1</sub>-6alkylthio, OH, CN, CF<sub>3</sub>, and CO<sub>2</sub>R<sup>8</sup>;

R<sup>8</sup> is selected from the group consisting of

- (a) hydrogen,  
(b) C<sub>1</sub>-6alkyl; and

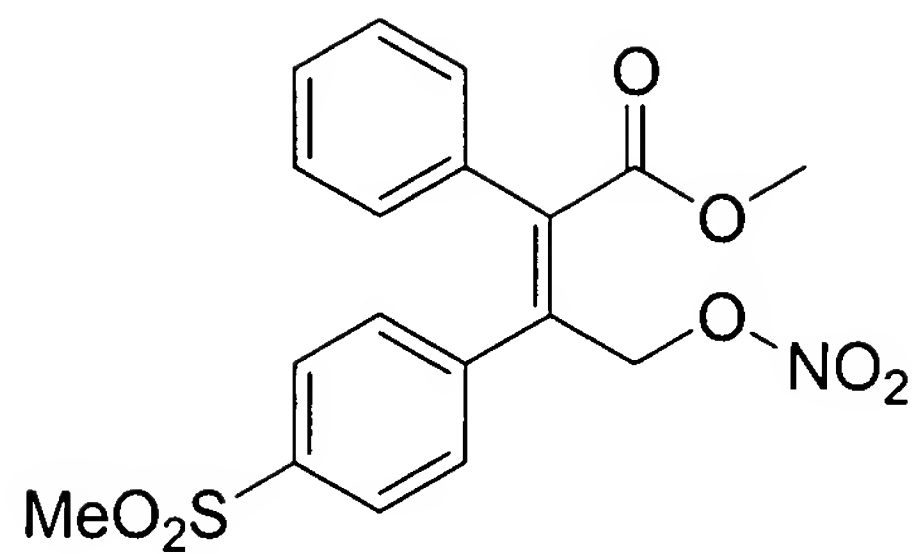
HET<sup>1</sup>, HET<sup>2</sup>, HET<sup>4</sup> and HET<sup>5</sup> are each independently selected from the group consisting of: benzimidazolyl, benzofuranyl, benzopyrazolyl, benzotriazolyl, benzothiophenyl, benzoxazolyl, carbazolyl, carbolinyl, cinnolinyl, furanyl, imidazolyl, indolinyl, indolyl, indolazinyl, indazolyl, isobenzofuranyl, isoindolyl, isoquinolyl, isothiazolyl, isoxazolyl, naphthyridinyl, oxadiazolyl, oxazolyl, pyrazinyl, pyrazolyl, pyridopyridinyl, pyridazinyl, pyridyl, pyrimidyl, pyrrolyl, quinazolinyl, quinolyl, quinoxalinyl, thiadiazolyl, thiazolyl, thienyl, triazolyl, azetidyl, 1,4-dioxanyl, hexahydroazepinyl, piperazinyl, piperidinyl, pyrrolidinyl, morpholinyl, thiomorpholinyl, dihydrobenzimidazolyl, dihydrobenzofuranyl, dihydrobenzothiophenyl, dihydrobenzoxazolyl, dihydrofuranyl, dihydroimidazolyl, dihydroindolyl, dihydroisooxazolyl, dihydroisothiazolyl, dihydrooxadiazolyl, dihydrooxazolyl, dihydropyrazinyl, dihydropyrazolyl, dihydropyridinyl, dihydropyrimidinyl, dihydropyrrolyl, dihydroquinolinyl, dihydrotetrazolyl, dihydrothiadiazolyl, dihydrothiazolyl, dihydrothienyl, dihydrotriazolyl, dihydroazetidyl, methylenedioxybenzoyl, tetrahydrofuranyl, and tetrahydrothienyl.

22.-25. (canceled)

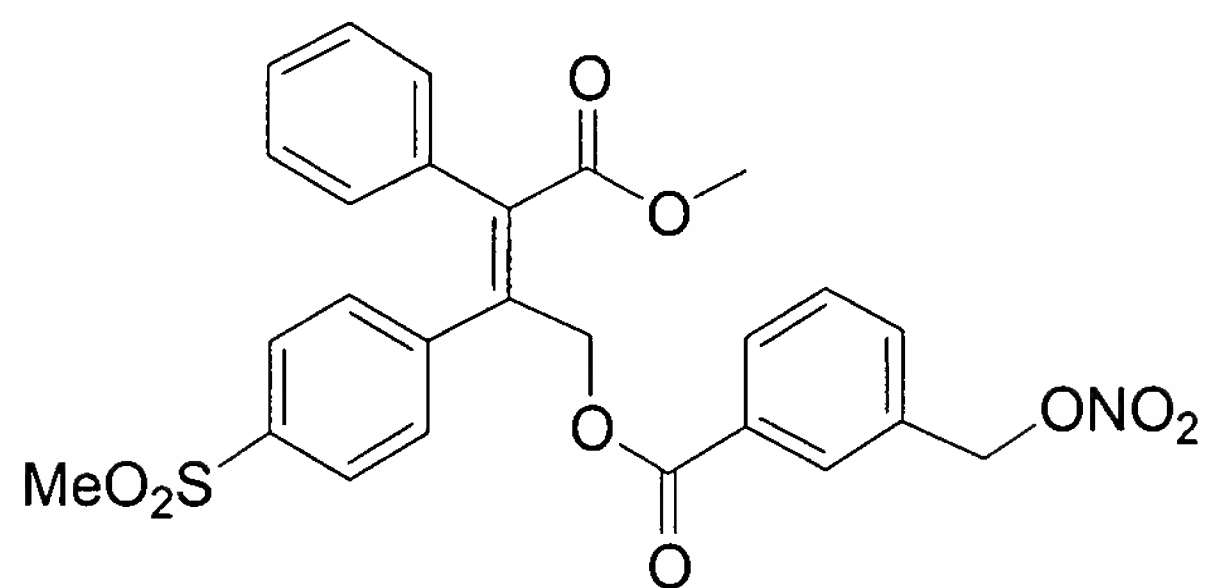
26. The compound according to Claim 1 wherein: R<sup>4</sup> is C<sub>1</sub>-6alkyl, mono-substituted with

- (i) N(R<sup>i</sup>)R<sup>ii</sup>, wherein R<sup>i</sup> and R<sup>ii</sup> are each independently selected from the group consisting of hydrogen and C<sub>1</sub>-4alkyl or  
(ii) -CO<sub>2</sub>R<sup>iii</sup>, wherein R<sup>iii</sup> is hydrogen or C<sub>1</sub>-4alkyl.

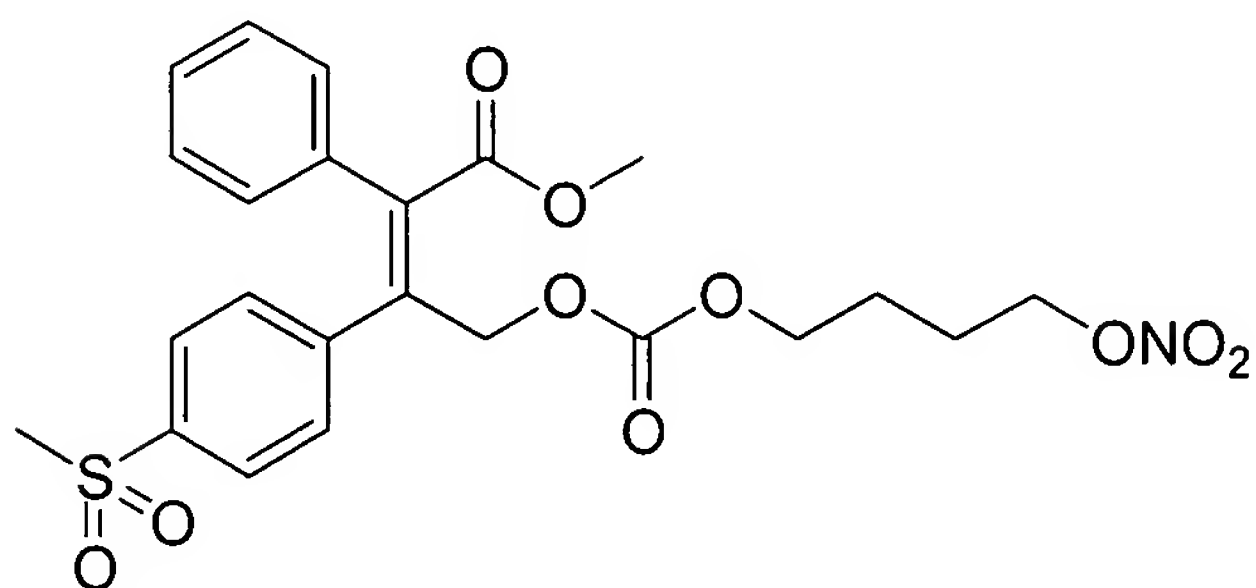
27. A compound selected from the following group:



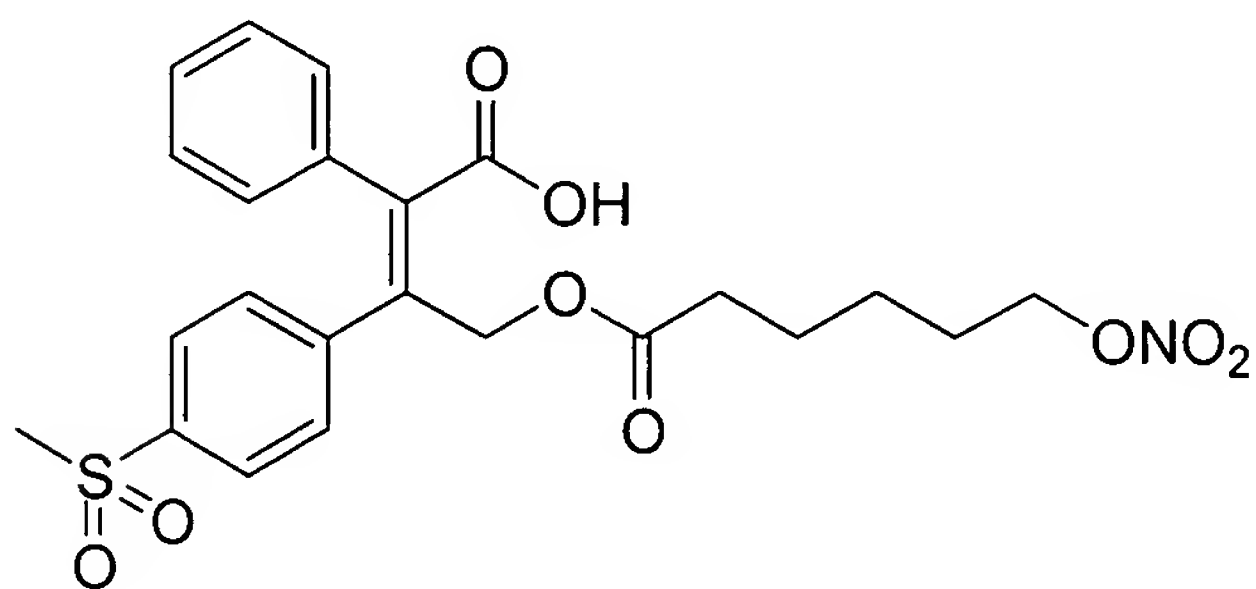
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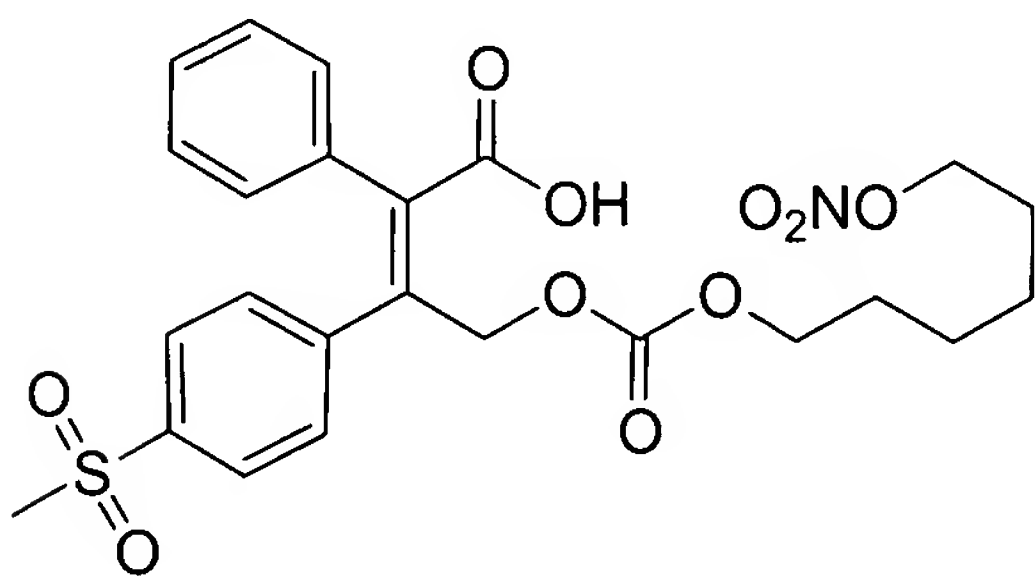
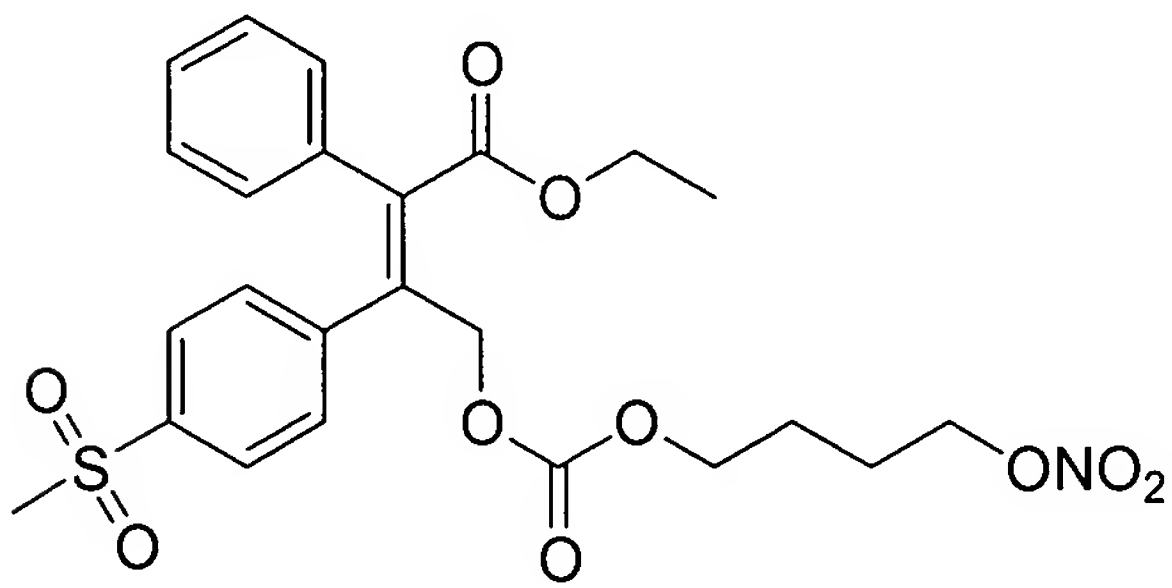
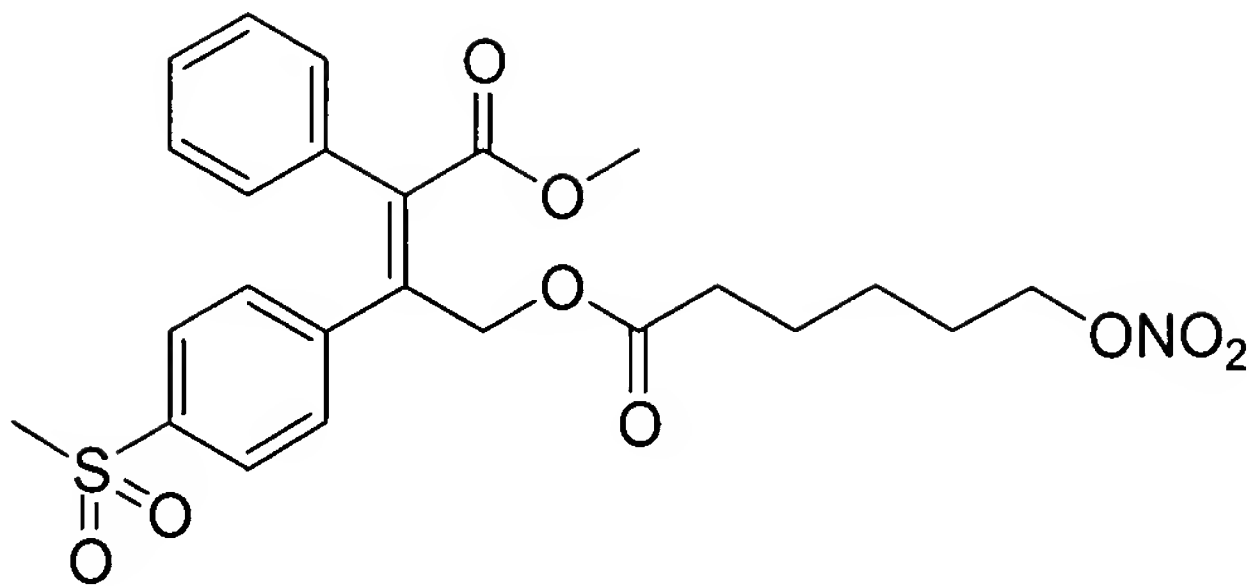


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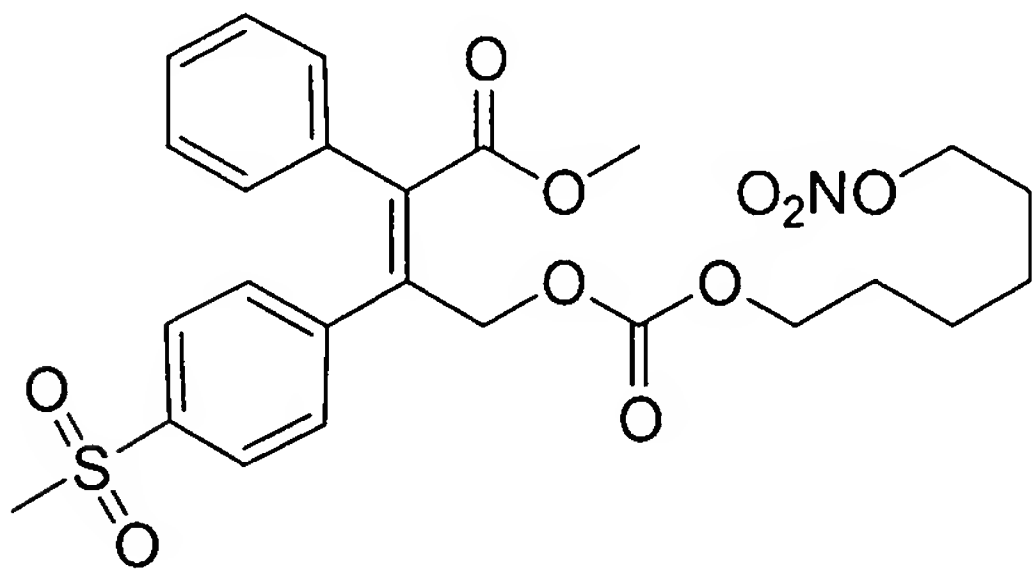


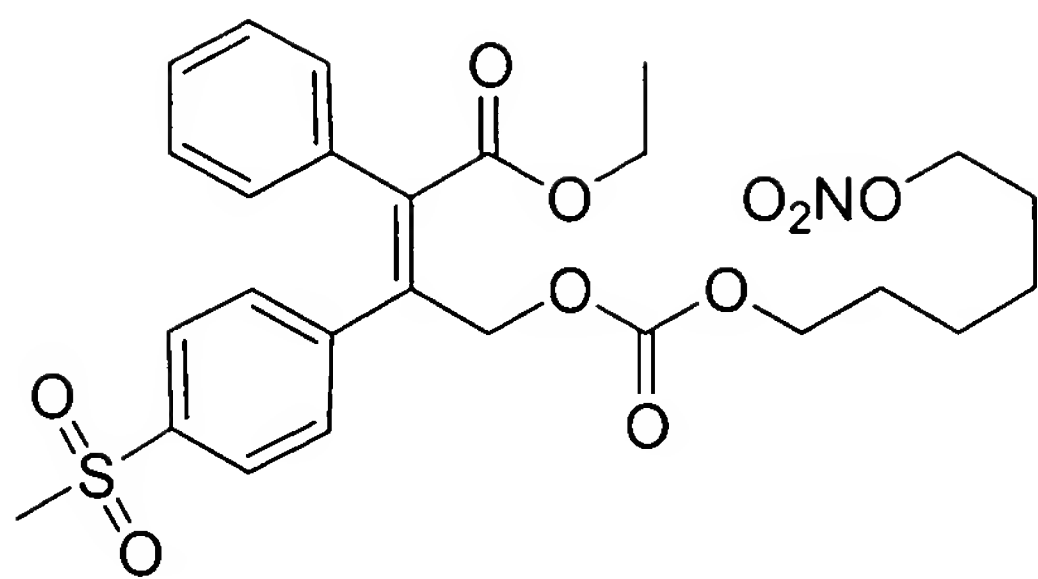
thereof,

or a pharmaceutically acceptable salt

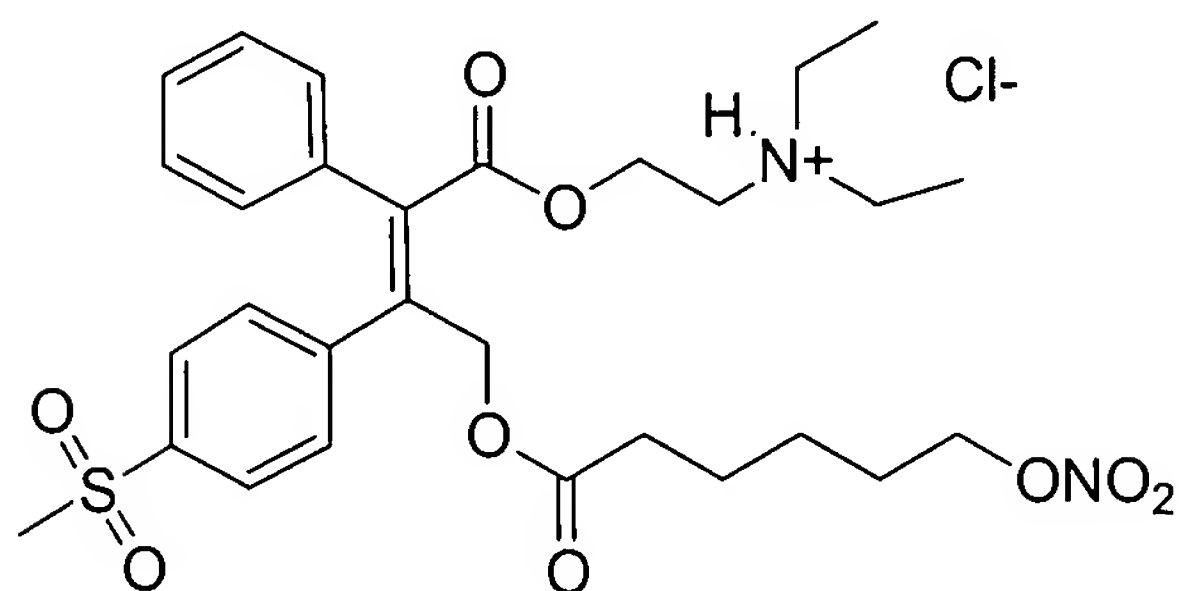


or a pharmaceutically acceptable salt thereof,



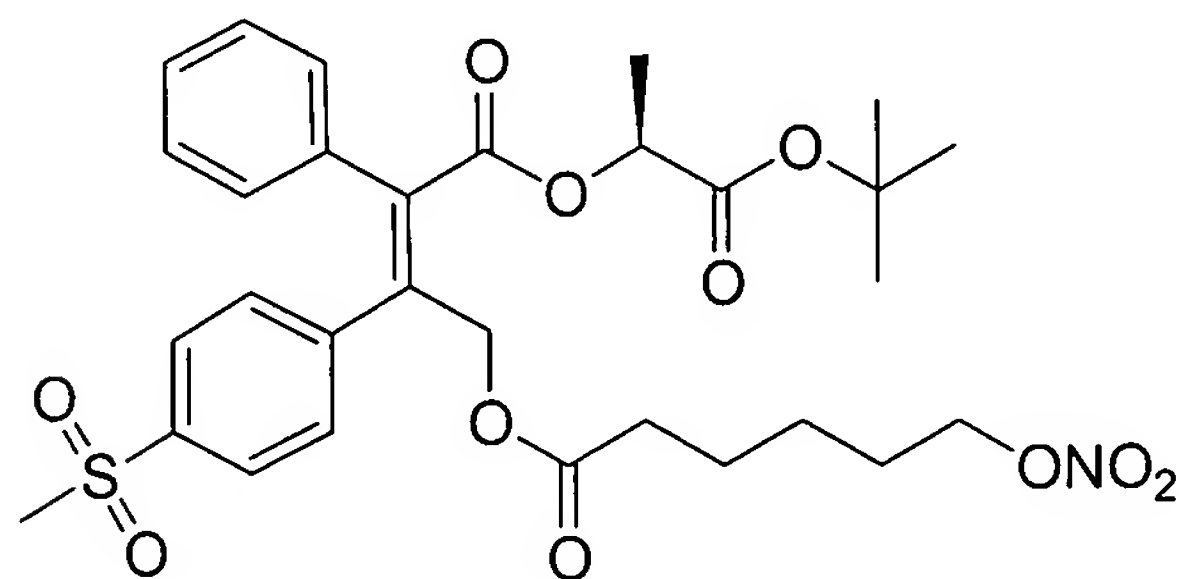


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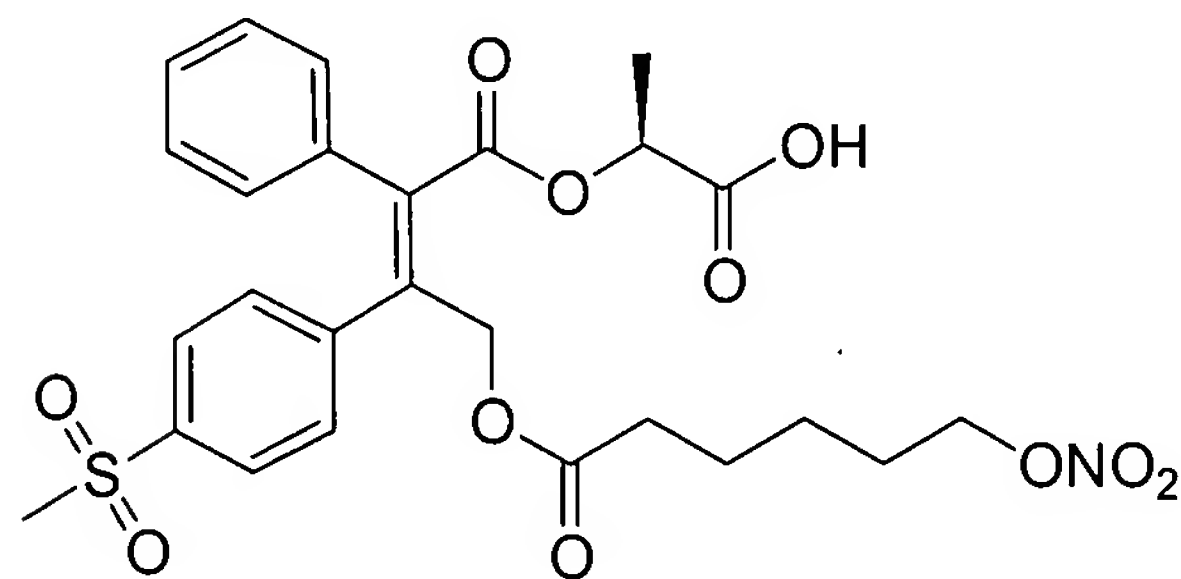


or a pharmaceutically acceptable salt

thereof,

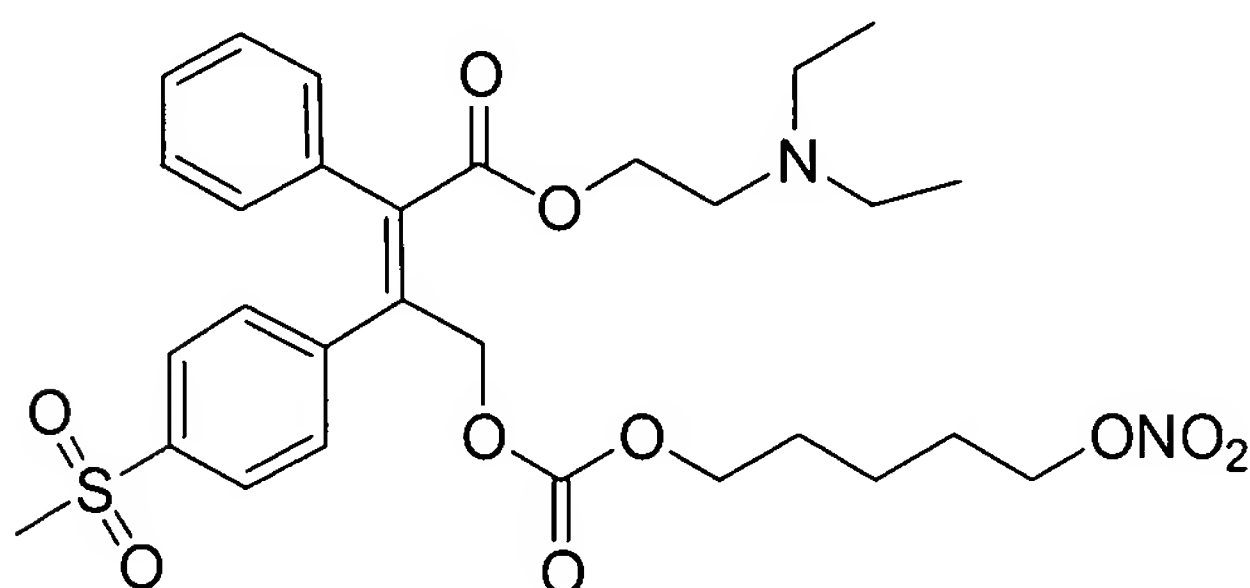
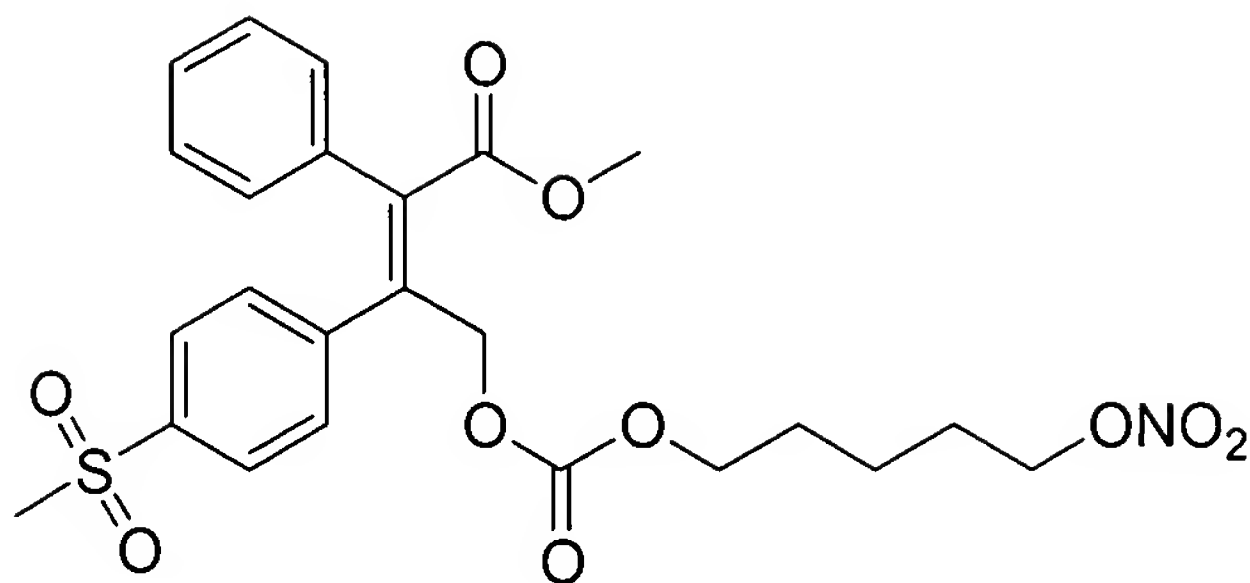


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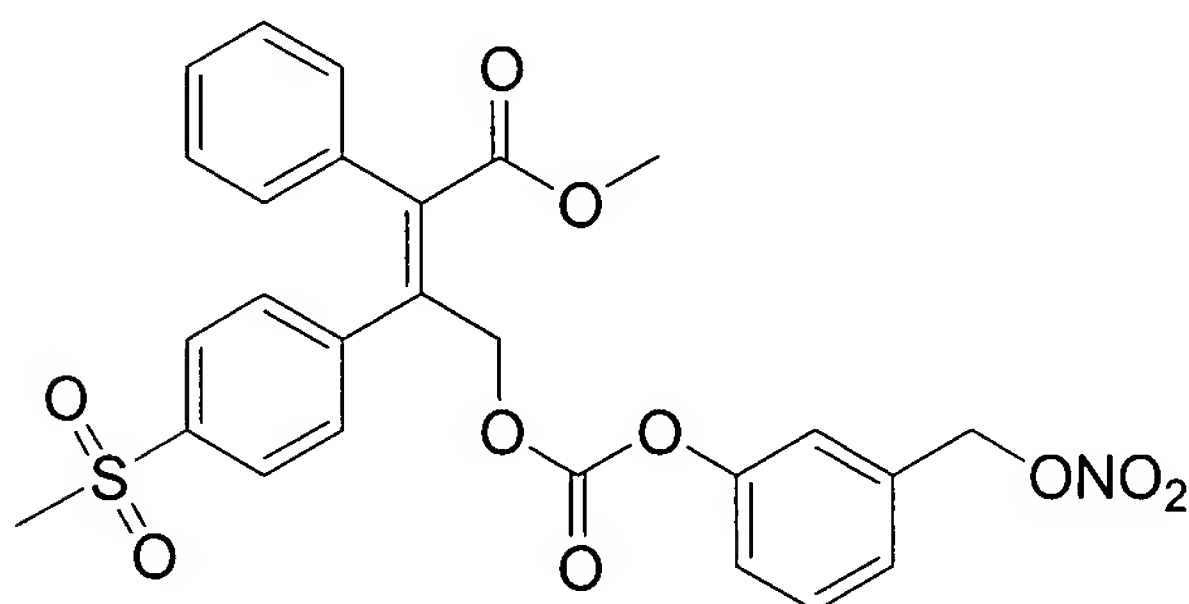
or a pharmaceutically acceptable salt

thereof,



or a pharmaceutically acceptable salt

thereof, and



28.-31. (canceled)

32. A method for treating a chronic cyclooxygenase-2 mediated disease or condition and reducing the risk of a thrombotic cardiovascular event in a human patient in need of such treatment and at risk of a thrombotic cardiovascular event comprising orally concomitantly or sequentially administering to said patient a compound according to Claim 1 in an amount effective to treat the cyclooxygenase-2 mediated disease or condition and aspirin in an amount effective to reduce the risk of the thrombotic cardiovascular event.

33.-40. (canceled)

41. (currently amended) A pharmaceutical composition comprising a compound of formula I according to ~~any one of claims~~ claim 1 ~~to 27~~, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier.

42. – 44. (canceled)